```
=> d his

     (FILE 'HOME' ENTERED AT 11:19:10 ON 07 FEB 2004)

     FILE 'CAPLUS' ENTERED AT 11:19:19 ON 07 FEB 2004
L1             2 S WO9901421/PN
               SELECT  L1 1 RN
L2         16289 S E1-E20

     FILE 'REGISTRY' ENTERED AT 11:20:02 ON 07 FEB 2004
L3             1 S 197520-71-1/RN
               SET NOTICE 1 DISPLAY
               SET NOTICE LOGIN DISPLAY

     FILE 'REGISTRY' ENTERED AT 11:20:20 ON 07 FEB 2004
L4             1 S 219796-67-5/RN
               SET NOTICE 1 DISPLAY
               SET NOTICE LOGIN DISPLAY
L5            11 S E4-E14

     FILE 'CAPLUS' ENTERED AT 11:21:46 ON 07 FEB 2004
L6             6 S L5

     FILE 'EUROPATFULL, PATDPAFULL, PCTFULL, RDISCLOSURE, USPATFULL, USPAT2'
     ENTERED AT 11:29:13 ON 07 FEB 2004
L7            18 S MEK(5A)INHIBITORS AND (WARNER(3A)LAMBERT OR PFIZER)

     FILE 'CAPLUS' ENTERED AT 11:32:43 ON 07 FEB 2004
L8             3 S L6 AND PAIN
```

10/049,762

Brain Research
Volume 566, Issues 1-2 , 6 December 1991, Pages 95-102

doi:10.1016/0006-8993(91)91685-T Cite or link using doi

Research report

Neurochemical studies on the mesolimbic circuitry of antinociception

Q. P. Ma and J. S. Han

Neuroscience Research Center, Beijing Medical University, Beijing, People's Republic of China

Accepted 16 July 1991. ; Available online 7 March 2003.

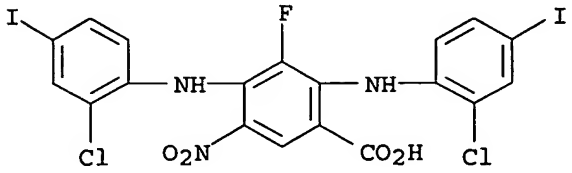
Abstract

Previous studies using the technique of microinjection into brain nuclei indicated that the periaqueductal gray (PAG), nucleus accumbens, habenula and amygdala play an essential role in pain modulation and that these nuclei possibly act through a `mesolimbic neural loop` to exert an analgesic effect, in which Met-enkephalin (MEK) and β-endorphin (β-EP) have been implicated as the two major opioid peptides involved in antinociception. In the present study performed in rabbits, intracranial microinjection was supplemented with push-pull perfusion and radioimmunoassay to determine whether the release of enkephalins (ENK) and β-EP was increased in these nuclei when the putative neural circuit was activated by morphine administered into one of the nuclei. The results showed: (1) microinjection of morphine into the PAG increased the release of ENK and β-EP in the N. accumbens, and vice versa; (2) microinjection of morphine into the N. accumbens increased the release of ENK and β-EP in the amygdala, and vice versa; (3) morphine microinjected into the PAG caused an increase in the release of ENK and β-EP in the amygdala and vice versa, although the release of ENK in PAG was statistically not significant. These results indicate that PAG, N. accumbens and amygdala are connected in a network served by a positive feedback circuitry.

Author Keywords: Periaqueductal gray; Nucleus accumbens; Amygdala; Morphine; Enkephalin; β-Endorphin

h c e ce ec c c e ce b c e e

L4 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2004 ACS on STN
RN **219796-67-5** REGISTRY
CN Benzoic acid, 2,4-bis[(2-chloro-4-iodophenyl)amino]-3-fluoro-5-nitro-
    (9CI) (CA INDEX NAME)
OTHER NAMES:
CN 2,4-Bis(2-chloro-4-iodophenylamino)-3-fluoro-5-nitrobenzoic acid
FS 3D CONCORD
MF C19 H10 Cl2 F I2 N3 O4
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

6 REFERENCES IN FILE CA (1907 TO DATE)
6 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> SET NOTICE LOGIN DISPLAY

NOTICE SET TO OFF FOR DISPLAY COMMAND
SET COMMAND COMPLETED

```
=> s wo9901421/pn
L1             2 WO9901421/PN

=> select  l1
ENTER ANSWER NUMBER OR RANGE (1-):1
ENTER DISPLAY CODE (TI) OR ?:rn
E1 THROUGH E20 ASSIGNED

=> s e1-e20
          1787 114977-28-5/BI
            32 162652-95-1/BI
            14 197520-71-1/BI
             6 219796-67-5/BI G4
             3 284018-87-7/BI
             3 284018-90-2/BI
             3 284018-93-5/BI
             3 284018-96-8/BI
             3 284018-99-1/BI
             3 284019-03-0/BI
             3 284019-06-3/BI
             3 284019-08-5/BI
             2 321167-78-6/BI
             2 321167-81-1/BI E14
          8032 33069-62-4/BI
            27 42016-93-3/BI
          5246 57-22-7/BI
            49 61079-72-9/BI
           787 71486-22-1/BI
          4368 865-21-4/BI
L2       16289 (114977-28-5/BI OR 162652-95-1/BI OR 197520-71-1/BI OR 219796-67
               -5/BI OR 284018-87-7/BI OR 284018-90-2/BI OR 284018-93-5/BI OR
               284018-96-8/BI OR 284018-99-1/BI OR 284019-03-0/BI OR 284019-06-
               3/BI OR 284019-08-5/BI OR 321167-78-6/BI OR 321167-81-1/BI OR
               33069-62-4/BI OR 42016-93-3/BI OR 57-22-7/BI OR 61079-72-9/BI
               OR 71486-22-1/BI OR 865-21-4/BI)

=> FIL REGISTRY
```

```
=> s l5
L6            6 L5

=> d ibib 1-6

L6   ANSWER 1 OF 6  CAPLUS  COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER:         2003:35363  CAPLUS
DOCUMENT NUMBER:          138:89582
TITLE:                    Preparation of 4-arylamino, 4-aryloxy, and 4-arylthio
                          diarylamines and derivatives as selective MEK
                          inhibitors for use as immunomodulators,
                          anti-inflammatory agents, and antiproliferative agents
INVENTOR(S):              Barrett, Stephen; Tecle, Haile
PATENT ASSIGNEE(S):       Warner-Lambert Company, USA
SOURCE:                   U.S., 16 pp., Cont.-in-part of U.S. Ser. No. 462,319.
                          CODEN: USXXAM
DOCUMENT TYPE:            Patent
LANGUAGE:                 English
FAMILY ACC. NUM. COUNT:   3
PATENT INFORMATION:

     PATENT NO.        KIND  DATE           APPLICATION NO.  DATE
     ---------------   ----  --------       ---------------  --------
     US 6506798        B1    20030114       US 2001-889084   20010711
     WO 9901421        A1    19990114       WO 1998-US13105  19980624
         W:  AL, AU, BA, BB, BG, BR, CA, CN, CZ, EE, GE, GW, HU, ID, IL, IS,
             JP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG,
             SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD,
             RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
             FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
             CM, GA, GN, ML, MR, NE, SN, TD, TG
     WO 2000041994     A1    20000720       WO 1999-US30418  19991221
         W:  AE, AL, AU, BA, BB, BG, BR, CA, CN, CR, CU, CZ, DM, EE, GD, GE,
             HR, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MA, MG,
             MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ,
             VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
             DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
             CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     US 6310060        B1    20011030       US 2000-462319   20000105
PRIORITY APPLN. INFO.:                    US 1997-51433P   P  19970701
                                          WO 1998-US13105  W  19980624
                                          WO 1999-US30418  W  19991221
                                          US 2000-462319   A2 20000105
                                          US 1999-115670P  P  19990113
                                          US 1999-122421P  P  19990302
OTHER SOURCE(S):          MARPAT 138:89582
REFERENCE COUNT:          10    THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS
                                RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6   ANSWER 2 OF 6  CAPLUS  COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER:         2001:63820  CAPLUS
DOCUMENT NUMBER:          134:131318
TITLE:                    Preparation of (phenylamino)benzenesulfonamides and
                          (phenylamino)benzamides as MEK inhibitors for the
                          treatment of chronic pain
INVENTOR(S):              Bridges, Alexander James; Booth, Richard John; Tecle,
                          Haile; Scaggs, Yvonne; Kaufman, Michael; Barrett,
                          Stephen Douglas; Dixon, Alistair; Lee, Kevin; Pinnock,
                          Robert Denham
PATENT ASSIGNEE(S):       Warner-Lambert Company, USA
SOURCE:                   PCT Int. Appl., 158 pp.
                          CODEN: PIXXD2
```

DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001005393	A2	20010125	WO 2000-US18348	20000705
WO 2001005393	A3	20010510		
W: AE, AG, AL, AU, BA, BB, BG, BR, BZ, CA, CN, CR, CU, CZ, DM, DZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MA, MG, MK, MN, MX, MZ, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1202724	A2	20020508	EP 2000-945140	20000705
EP 1202724	B1	20031001		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
AT 250932	E	20031015	AT 2000-945140	20000705
ZA 2001009909	A	20030228	ZA 2001-9909	20011130

PRIORITY APPLN. INFO.:
US 1999-144280P P 19990716
US 1999-144320P P 19990716
US 1999-144419P P 19990716
US 1999-144655P P 19990716
US 1999-144658P P 19990716
US 1999-144659P P 19990716
WO 2000-US18348 W 20000705

OTHER SOURCE(S): MARPAT 134:131318

L6 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER: 2001:63819 CAPLUS
DOCUMENT NUMBER: 134:131317
TITLE: Preparation of 2-phenylaminobenzamides and analogs as MEK inhibitors for the treatment of chronic pain
INVENTOR(S): Dixon, Alistair; Lee, Kevin; Pinnock, Robert Denham
PATENT ASSIGNEE(S): Warner-Lambert Company, USA
SOURCE: PCT Int. Appl., 132 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001005392	A2	20010125	WO 2000-US18347	20000705
WO 2001005392	A3	20010719		
W: AE, AG, AL, AU, BA, BB, BG, BR, BZ, CA, CN, CR, CU, CZ, DM, DZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MA, MG, MK, MN, MX, MZ, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1202726	A2	20020508	EP 2000-943383	20000705
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
ZA 2001009907	A	20030228	ZA 2001-9907	20011130

PRIORITY APPLN. INFO.:
US 1999-144292P P 19990716
WO 2000-US18347 W 20000705

OTHER SOURCE(S): MARPAT 134:131317

L6 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER: 2001:63818 CAPLUS
DOCUMENT NUMBER: 134:131540
TITLE: Preparation of (2-heterocyclylphenyl)(4-iodophenyl)amines as MEK inhibitors for the treatment of chronic pain
INVENTOR(S): Barrett, Stephen Douglas; Bridges, Alexander James; Tecle, Haile; Dixon, Alistair; Lee, Kevin; Pinnock, Robert Denham; Zhang, Lu-Yan
PATENT ASSIGNEE(S): Warner-Lambert Company, USA
SOURCE: PCT Int. Appl., 104 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001005391	A2	20010125	WO 2000-US18346	20000705
WO 2001005391	A3	20010719		
W: AE, AG, AL, AU, BA, BB, BG, BR, BZ, CA, CN, CR, CU, CZ, DM, DZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MA, MG, MK, MN, MX, MZ, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1202732	A2	20020508	EP 2000-943382	20000705
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
ZA 2001009903	A	20030228	ZA 2001-9903	20011130

PRIORITY APPLN. INFO.: US 1999-144403P P 19990716
WO 2000-US18346 W 20000705
OTHER SOURCE(S): MARPAT 134:131540

L6 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER: 2000:493498 CAPLUS
DOCUMENT NUMBER: 133:104875
TITLE: Preparation of N-iodophenylanthranilates and analogs as MEK inhibitors
INVENTOR(S): Barrett, Stephen Douglas; Tecle, Haile
PATENT ASSIGNEE(S): Warner-Lambert Company, USA
SOURCE: PCT Int. Appl., 49 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 3
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000041994	A1	20000720	WO 1999-US30418	19991221
W: AE, AL, AU, BA, BB, BG, BR, CA, CN, CR, CU, CZ, DM, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MA, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
JP 2000204068	A2	20000725	JP 1999-53560	19990302
EP 1144362	A1	20011017	EP 1999-968152	19991221
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				

BR 9916798	A	20011023	BR 1999-16798		19991221
JP 2002534491	T2	20021015	JP 2000-593563		19991221
US 6506798	B1	20030114	US 2001-889084		20010711

PRIORITY APPLN. INFO.:

US 1999-115670P	P	19990113
US 1999-122421P	P	19990302
US 1997-51433P	P	19970701
WO 1998-US13105	W	19980624
WO 1999-US30418	W	19991221
US 2000-462319	A2	20000105

OTHER SOURCE(S): MARPAT 133:104875

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1999:48694 CAPLUS

DOCUMENT NUMBER: 130:124898

TITLE: Preparation of 2-(4-bromo or 4-iodo phenylamino)benzoic acid derivatives as MEK inhibitors

INVENTOR(S): Barrett, Stephen Douglas; Bridges, Alexander James; Cody, Donna Reynolds; Doherty, Annette Marian; Dudley, David Thomas; Saltiel, Alan Robert; Schroeder, Mel Conrad; Tecle, Haile

PATENT ASSIGNEE(S): Warner-Lambert Company, USA

SOURCE: PCT Int. Appl., 67 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9901421	A1	19990114	WO 1998-US13105	19980624
W: AL, AU, BA, BB, BG, BR, CA, CN, CZ, EE, GE, GW, HU, ID, IL, IS, JP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
AU 9882626	A1	19990125	AU 1998-82626	19980624
AU 756586	B2	20030116		
EP 993437	A1	20000419	EP 1998-932829	19980624
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO				
BR 9810385	A	20000905	BR 1998-10385	19980624
JP 2002509536	T2	20020326	JP 1999-507227	19980624
NZ 501277	A	20021220	NZ 1998-501277	19980624
ZA 9805726	A	19990127	ZA 1998-5726	19980630
MX 9910556	A	20000430	MX 1999-10556	19991116
US 6310060	B1	20011030	US 2000-462319	20000105
US 6506798	B1	20030114	US 2001-889084	20010711
US 2002022647	A1	20020221	US 2001-931596	20010816
US 6492363	B2	20021210		
US 2003149015	A1	20030807	US 2002-315654	20021210

PRIORITY APPLN. INFO.:

US 1997-51433P	P	19970701
WO 1998-US13105	W	19980624
WO 1999-US30418	W	19991221
US 2000-462319	A2	20000105
US 2001-931596	A3	20010816

OTHER SOURCE(S): MARPAT 130:124898

REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
=> s l6 and pain
         31759 PAIN
           833 PAINS
         32361 PAIN
                 (PAIN OR PAINS)
L8            3 L6 AND PAIN

=> d ibib 1-3

L8   ANSWER 1 OF 3  CAPLUS  COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER:         2001:63820  CAPLUS
DOCUMENT NUMBER:          134:131318
TITLE:                    Preparation of (phenylamino)benzenesulfonamides and
                          (phenylamino)benzamides as MEK inhibitors for the
                          treatment of chronic pain
INVENTOR(S):              Bridges, Alexander James; Booth, Richard John; Tecle,
                          Haile; Scaggs, Yvonne; Kaufman, Michael; Barrett,
                          Stephen Douglas; Dixon, Alistair; Lee, Kevin; Pinnock,
                          Robert Denham
PATENT ASSIGNEE(S):       Warner-Lambert Company, USA
SOURCE:                   PCT Int. Appl., 158 pp.
                          CODEN: PIXXD2
DOCUMENT TYPE:            Patent
LANGUAGE:                 English
FAMILY ACC. NUM. COUNT:   1
PATENT INFORMATION:

     PATENT NO.        KIND  DATE           APPLICATION NO.  DATE
     ---------------   ----  --------       ---------------  --------
     WO 2001005393      A2   20010125       WO 2000-US18348  20000705
     WO 2001005393      A3   20010510
         W:  AE, AG, AL, AU, BA, BB, BG, BR, BZ, CA, CN, CR, CU, CZ, DM, DZ,
             EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT,
             LV, MA, MG, MK, MN, MX, MZ, NO, NZ, PL, RO, SG, SI, SK, SL, TR,
             TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
             CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     EP 1202724         A2   20020508       EP 2000-945140   20000705
     EP 1202724         B1   20031001
         R:  AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL
     AT 250932          E    20031015       AT 2000-945140   20000705
     ZA 2001009909      A    20030228       ZA 2001-9909     20011130
PRIORITY APPLN. INFO.:                   US 1999-144280P  P  19990716
                                         US 1999-144320P  P  19990716
                                         US 1999-144419P  P  19990716
                                         US 1999-144655P  P  19990716
                                         US 1999-144658P  P  19990716
                                         US 1999-144659P  P  19990716
                                         WO 2000-US18348  W  20000705
OTHER SOURCE(S):          MARPAT 134:131318

L8   ANSWER 2 OF 3  CAPLUS  COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER:         2001:63819  CAPLUS
DOCUMENT NUMBER:          134:131317
TITLE:                    Preparation of 2-phenylaminobenzamides and analogs as
                          MEK inhibitors for the treatment of chronic
                          pain
INVENTOR(S):              Dixon, Alistair; Lee, Kevin; Pinnock, Robert Denham
PATENT ASSIGNEE(S):       Warner-Lambert Company, USA
SOURCE:                   PCT Int. Appl., 132 pp.
                          CODEN: PIXXD2
```

DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001005392	A2	20010125	WO 2000-US18347	20000705
WO 2001005392	A3	20010719		
W: AE, AG, AL, AU, BA, BB, BG, BR, BZ, CA, CN, CR, CU, CZ, DM, DZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MA, MG, MK, MN, MX, MZ, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1202726	A2	20020508	EP 2000-943383	20000705
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
ZA 2001009907	A	20030228	ZA 2001-9907	20011130

PRIORITY APPLN. INFO.: US 1999-144292P P 19990716
WO 2000-US18347 W 20000705
OTHER SOURCE(S): MARPAT 134:131317

L8 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER: 2001:63818 CAPLUS
DOCUMENT NUMBER: 134:131540
TITLE: Preparation of (2-heterocyclylphenyl)(4-iodophenyl)amines as MEK inhibitors for the treatment of chronic **pain**
INVENTOR(S): Barrett, Stephen Douglas; Bridges, Alexander James; Tecle, Haile; Dixon, Alistair; Lee, Kevin; Pinnock, Robert Denham; Zhang, Lu-Yan
PATENT ASSIGNEE(S): Warner-Lambert Company, USA
SOURCE: PCT Int. Appl., 104 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001005391	A2	20010125	WO 2000-US18346	20000705
WO 2001005391	A3	20010719		
W: AE, AG, AL, AU, BA, BB, BG, BR, BZ, CA, CN, CR, CU, CZ, DM, DZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MA, MG, MK, MN, MX, MZ, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
EP 1202732	A2	20020508	EP 2000-943382	20000705
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
ZA 2001009903	A	20030228	ZA 2001-9903	20011130

PRIORITY APPLN. INFO.: US 1999-144403P P 19990716
WO 2000-US18346 W 20000705
OTHER SOURCE(S): MARPAT 134:131540

10/049,762

L2 ANSWER 2 OF 2418 MEDLINE on STN
ACCESSION NUMBER: 2003419470 MEDLINE
DOCUMENT NUMBER: 22839652 PubMed ID: 12959289
TITLE: Analgesic efficacy of non-steroidal anti-**inflammatory** drugs in experimental **pain** in humans.
AUTHOR: Walker J S; Arroyo J F; Nguyen T; Day R O
CORPORATE SOURCE: Department of Clinical Pharmacology and Toxicology, Garvan Institute of Medical Research, St Vincent's Hospital, NSW 2010, Australia.
SOURCE: BRITISH JOURNAL OF CLINICAL PHARMACOLOGY, (1993 Nov) 36 (5) 417-25.
Journal code: 7503323. ISSN: 0306-5251.
PUB. COUNTRY: England: United Kingdom
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200309
ENTRY DATE: Entered STN: 20030909
Last Updated on STN: 20030918
Entered Medline: 20030917

AB 1. The aim of this study was to establish a simple and reliable experimental **pain** model that could distinguish the analgesic effects of non-steroidal anti-**inflammatory** drug (NSAID) treatment from placebo in human volunteers. 2. The reproducibility and reliability over time of subject pain ratings was compared using cutaneous electrical stimuli applied to either the thenar eminence or the ear lobe at varying intensities and modes. Subjects were asked to respond firstly, when the stimulus became clearly sharp and painful ('first pain') and secondly, when the sensation became deep and burning and no further increase in stimulus intensity could be tolerated ('second pain'). 3. Constant voltage stimuli were found to be more reproducible than constant current stimuli. Both phasic (intermittent) and tonic (continuous) stimulation modalities produced 'first' and 'second pain' sensations. The latter sensation was more reproducible, and was perceived as a burning pain which is akin to clinical pain. 4. Analgesics from the NSAID class were found to attenuate reliably only 'second pain' sensations. The analgesic effects of ibuprofen (ibuprofen vs placebo: 0.12 +/- 0.09 vs 0.02 +/- 0.07 volt h(-1), P = 0.03; 95% confidence interval for differences (CI): 0.03-0.18) and diflunisal (diflunisal vs placebo: 0.29 +/- 0.40 vs 0.005 +/- 0.27 volt h(-1), P = 0.0001; CI: 0.168-0.407), respectively, could be distinguished from placebo.

TI Analgesic efficacy of non-steroidal anti-**inflammatory** drugs in experimental **pain** in humans.

AB 1. The aim of this study was to establish a simple and reliable experimental **pain** model that could distinguish the analgesic effects of non-steroidal anti-**inflammatory** drug (NSAID) treatment from placebo in human volunteers. 2. The reproducibility and reliability over time of subject pain ratings was. . .

=>